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| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE | TOTAL |
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L50 0 L49

Text search

=> fil medl,biosis,hcap,embase,jicst,wpids;s ~~tr~~ methane and (cellular prolifer? or inflam? disease or fibro? or lymphocy? or glomerulonephri?)

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| COST IN U.S. DOLLARS | SINCE FILE | TOTAL |
| | ENTRY | SESSION |
| FULL ESTIMATED COST | 0.76 | 929.23 |

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| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE | TOTAL |
| | ENTRY | SESSION |
| CA SUBSCRIBER PRICE | 0.00 | -10.62 |

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L51 0 FILE MEDLINE
 L52 0 FILE BIOSIS
 L53 0 FILE HCAPLUS
 L54 0 FILE EMBASE

Searched by: Mary Hale 308-4258 CM-1 1E01

L55 0 FILE JICST-EPLUS
L56 0 FILE WPIDS

TOTAL FOR ALL FILES

L57 0 TRIARY METHANE AND (CELLULAR PROLIFER? OR INFLAM? DISEASE OR
FIBRO? OR LYMPHOCY? OR GLOMERULONEPHRI?)

=> s (triary? or tricycl?) (1) methane and (cellular prolifer? or inflam? disease or
fibro? or lymphocy? or glomerulonephri?)

L58 1 FILE MEDLINE
L59 1 FILE BIOSIS
L60 1 FILE HCAPLUS
L61 1 FILE EMBASE
L62 0 FILE JICST-EPLUS
L63 5 FILE WPIDS

TOTAL FOR ALL FILES

L64 9 (TRIARY? OR TRICYCL?) (L) METHANE AND (CELLULAR PROLIFER? OR
INFLAM? DISEASE OR FIBRO? OR LYMPHOCY? OR GLOMERULONEPHRI?)

=> s brugnara, c?/au,in

'IN' IS NOT A VALID FIELD CODE

L65 129 FILE MEDLINE
L66 235 FILE BIOSIS
L67 118 FILE HCAPLUS
'IN' IS NOT A VALID FIELD CODE
L68 129 FILE EMBASE
L69 0 FILE JICST-EPLUS
L70 18 FILE WPIDS

TOTAL FOR ALL FILES

L71 629 BRUGNARA, C?/AU,IN

=> dup rem l64

PROCESSING COMPLETED FOR L64

L72 6 DUP REM L64 (3 DUPLICATES REMOVED)

=> d cbib abs 1-6

L72 ANSWER 1 OF 6 WPIDS (C) 2002 THOMSON DERWENT

AN 2002-411969 [44] WPIDS

AB JP2002114837 A UPAB: 20020711

NOVELTY - A new epoxy resin composition comprises an epoxy resin-(A) having at least three epoxy groups, a curing agent-(B) of specific formula, a fire retardant-(C), and spherical fused silica-(D).

DETAILED DESCRIPTION - A new epoxy resin composition comprises an epoxy resin-(A) having at least three epoxy groups, a curing agent-(B) of formula (1-1), a fire retardant-(C), and spherical fused silica-(D).

Another composition further comprises a coupling agent-(E). Formula (1-1)-p

USE - For copper-plated laminated boards.

ADVANTAGE - The new composition provides copper-plated laminated circuit boards having improved heat resistance, lower thermal expansion, lower water absorptivity, and higher soldering resistance.
Dwg.0/0

L72 ANSWER 2 OF 6 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 1

2002:134482 Document No.: PREV200200134482. Use of **triaryl**

methane compounds for inhibiting unwanted **cellular proliferation** associated with **inflammatory**

disease. Brugnara, Carlo; Halperin, Jose; Bellot, Emile M., Jr. (1); Froimowitz, Mark; Lombardy, Richard John; Clifford, John J.; Gao, Ying-Duo; Haidar, Reem M.; Kelleher, Eugene W.; Kher, Falguni M.; Moussa, Adel M.; Sachdeva, Yesh P.; Sun, Minghua; Taft, Heather N.. (1) Beverly, MA USA. ASSIGNEE: Ion Pharmaceuticals, Inc., Cambridge, MA, USA. Patent Info.: US 6331564 December 18, 2001. Official Gazette of the United States Patent and Trademark Office Patents, (Dec. 18, 2001) Vol. 1253, No. 3, pp. No Pagination. <http://www.uspto.gov/web/menu/patdata.html>. e-file. ISSN: 0098-1133. Language: English.

AB The present invention provides a class of chemical compounds useful as efficacious drugs in the treatment of sickle cell disease and diseases characterized by unwanted or abnormal cell proliferation, and in particular **inflammatory diseases** associated with unwanted **cellular proliferation**. The active compounds are substituted **triaryl methane** compounds or analogues thereof where one or more of the aryl groups is replaced with a heteroaryl, cycloalkyl or heterocycloalkyl group and/or the tertiary carbon atom is replaced with a different atom such as Si, Ge, N or P, the compounds inhibit mammalian cell proliferation, inhibit the Gardos channel of erythrocytes, reduce sickle erythrocyte dehydration and/or delay the occurrence of erythrocyte sickling or deformation.

L72 ANSWER 3 OF 6 WPIDS (C) 2002 THOMSON DERWENT

AN 2001-514394 [56] WPIDS

AB WO 200149663 A UPAB: 20011001

NOVELTY - Method for treating or preventing autoimmune disorders, transplant rejection or graft versus-host disease by administering a **triaryl methane** compound (I), is new.

DETAILED DESCRIPTION - Method for treating or preventing autoimmune disorders, transplant rejection or graft versus-host disease by administering a **triaryl methane** compound of formula (I), is new.

An INDEPENDENT CLAIM is also included for suppressing antigen- or cytokine- or mitogen-stimulated calcium entry via store-operated calcium channels in **lymphocytes**, monocytes, macrophages, platelets and endothelial cells and/or cytokine production by these cells and/or activation of these cells of a patient, without concomitant cytochrome P450 inhibition, is new.

X, Y, Z = CH₂, O, S, NR₁, N=CH, CH=N or R₂-C=C-R₃;

R₂, R₃ = H or may together form a saturated or unsaturated carbocyclic or heterocyclic ring, optionally substituted with 1 or more R;

R₁ = H, alkyl, alkenyl, alkynyl, cycloalkyl, aryl, acyl or aroyl, optionally substituted with OH, optionally substituted amino, CN, alkoxy, halo, trihaloalkyl, NO₂, thio, alkylthio, carboxy or alkoxycarbonyl;

R = H, halo, trihaloalkyl, OH, acyloxy, alkoxy, alkenyloxy, thio, alkylthio, NO₂, CN, ureido, acyl, carboxy, alkoxycarbonyl, N(R₄)(R₅) or optionally unsaturated, chiral or achiral, cyclic or acyclic, straight or branched 1-20C hydrocarbonyl, optionally substituted with OH, halo, trihaloalkyl, alkylthio, alkoxy, carboxyl, alkoxycarbonyl, oxoalkyl, CN or N(R₄)(R₅);

R₄, R₅ = H, alkyl, alkenyl, alkynyl, cycloalkyl or acyl;

R₄+R₅ = a ring, where a C may be optionally substituted a heteroatom (O, S or NR₆);

R₆ = H, alkyl, alkenyl, alkynyl, cycloalkyl, hydroxyalkyl or carboxyalkyl;

n = 0-5;

m = 1-2; provided that when m = 1, Q = OH, CN, carboxyalkyl, N-(R₇)(R₈) or -NH-Het; and when m = 2, Q = a spacer of 2-10C either as a straight or branched hydrocarbon chain, or containing a hydrocarbon ring;

R₇, R₈ = H, 1-4C alkyl, cycloalkyl, aryl, acyl or amido;

R7+R8 = saturated or unsaturated heterocyclic ring optionally substituted with 1-3 additional N, O or S;
Het = thiazole, oxazole, isoxazole, pyridine, pyrimidine or purine;
U, V = H, O a group of formula (i).

An INDEPENDENT CLAIM is included for a method for inhibiting calcium activated potassium channel encoded by IKCa1 in a target cell type of a patient, without causing side effects due to concomitant inhibition of cytochrome P-450 enzyme activity, comprising administering a compound (I) that causes inhibition of calcium activated potassium channels encoded by IKCa1 in the target cell type of animals of the same species as the patient but which does not cause inhibition of activity of any cytochrome P-450 enzymes in any tissue of animals of the same species of the patient at concentrations at least 50 times greater than the half blocking concentration of that compound required for inhibition of the calcium activated potassium channels.

ACTIVITY - Immunosuppressive; antirheumatic; antiarthritic; dermatological; antiinflammatory; thyromimetic; neuroprotective; antidiabetic; nephrotropic; vasotropic; antipsoriatic; antipruritic; antiseborrheic; antiarteriosclerotic; ophthalmological; auditory; antianemic; osteopathic; antiulcer; antimigraine; antiallergic; hepatotropic; virucide.

MECHANISM OF ACTION - Calcium activated potassium channel inhibitor.

In a test to determine inhibition of the cloned human IKCa1 channel in COS-7 cells, 1-((2-chlorophenyl)diphenylmethyl)-1H pyrazole (Ia) had Kd 20 plus or minus 3 nM.

USE - For treating or preventing autoimmune disorders, transplant rejection or graft-versus-host disease (claimed), including rheumatoid arthritis, systemic lupus erythematosus, Hashimoto's thyroiditis, multiple sclerosis, myasthenia gravis, type I diabetes mellitus, nephrotic syndrome, steroid-dependent and steroid-resistant nephrosis, palmar-plantar pustulosis, allergic encephalomyelitis, **glomerulonephritis**, Behcet's syndrome, ankylosing spondylitis, polymyositis and **fibromyositis**; inflammatory, proliferative and hyperproliferative skin diseases, including psoriasis, psoriatic arthritis, atopic dermatitis, contact dermatitis, seborrheic dermatitis, Lichen planus, pemphigus, bullous pemphigus, epidermolysis bullosa, angiodemas, vasculitides, erythemas, cutaneous eosinophilias, acne, alopecia areata and arteriosclerosis. (I) are also useful for treating respiratory diseases, e.g. sarcoidosis, **fibroid** lung, idiopathic interstitial pneumonia and reversible obstructive airways disease, including asthma and bronchitis; hepatic injury associated with ischemia; and eye diseases, e.g. keratoconjunctivitis, vernal conjunctivitis, keratitis, uveitis, corneal leukoma, ocular pemphigus, Mooren's ulcer, scleritis, Graves' ophthalmopathy and sympathetic ophthalmia.

(I) may also be used to treat inflammatory bowel diseases (e.g. Crohn's disease), neurological disease (e.g. Guillain-Barre syndrome, Meniere's disease, radiculopathy), endocrine diseases (e.g. hyperthyroidism, Basedow's disease), hematological diseases (e.g. pure red cell aplasia, aplastic anemia, hypoplastic anemia, idiopathic thrombocytopenic purpura, autoimmune hemolytic anemia, agranulocytosis, anerythroplasia), bone diseases (e.g. osteoporosis), respiratory disease (e.g. sarcoidosis, idiopathic interstitial pneumonia), skin diseases (e.g. dermatomyositis, leukoderma vulgaris, ichthyosis vulgaris, photoallergic sensitivitiy, cutaneous T cell lymphoma), genitals (orchitis, vulvitis), circulatory diseases (e.g. arteriosclerosis, polyarteritis nodosa, vasculitis, Buerger's disease, myocardosis), collagen disorders (e.g. scleroderma, aortitis syndrome, eosinophilic fascitis, Wegener's granulomatosis, Sjogren's syndrome, periodontal diseases), kidney diseases (e.g. nephrotic syndrome, hemolytic-uremic syndrome, Goodpasture's

syndrome), and muscular dystrophy. Other diseases which can be treated include intestinal inflammation/allergies (e.g. coeliac disease, proctitis, ulcerative colitis, eosinophilic gastroenteritis, mastocytosis, Crohn's disease, ulcerative colitis), food-related allergic diseases which have symptomatic manifestations remote from the gastrointestinal tract, e.g. migraine, rhinitis and eczema.

(I) may be used for treating or preventing inflammation of mucosa or blood vessels, gastric ulcers, vascular damage caused by ischemic diseases and thrombosis, ischemic bowel diseases; also for treating multidrug resistance of tumor cells; and hepatic diseases, e.g. chronic autoimmune liver diseases including autoimmune hepatitis, primary biliary cirrhosis, sclerosing cholangitis, partial liver resection, acute liver necrosis, B-virus hepatitis, nonA/nonB hepatitis, cirrhosis.

(I) may be administered with other active agents, e.g. analgesics, antibiotics and other immunosuppressive drugs.

ADVANTAGE - (I) do not produce the side effects associated with currently available drugs.
Dwg.0/0

L72 ANSWER 4 OF 6 WPIDS (C) 2002 THOMSON DERWENT
AN 2001-183074 [18] WPIDS
CR 1997-011834 [01]; 2002-025885 [03]; 2002-443183 [47]
AB WO 200111077 A UPAB: 20020725

NOVELTY - Diagnosis of irritable bowel syndrome, **fibromyalgia**, chronic fatigue syndrome, depression, attention deficit/hyperactivity disorder, an autoimmune disease or Crohn's disease comprises detecting the presence of small intestinal bacterial overgrowth in a subject having at least one symptom associated with the diagnosis of one of these diseases.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(A) treatment of irritable bowel syndrome, **fibromyalgia**, chronic fatigue syndrome, depression, attention deficit/hyperactivity disorder, an autoimmune disease or Crohn's disease comprising detecting the presence of small intestinal bacterial overgrowth in a subject having at least one symptom associated with the diagnosis of one of these diseases, and eradicating the bacterial overgrowth; and

(B) a kit for the diagnosis of irritable bowel syndrome, **fibromyalgia**, chronic fatigue syndrome, depression, attention deficit/hyperactivity disorder, an autoimmune disease or Crohn's disease comprising at least one breath sampling container, a pre-measured amount of substrate and instructions for the user for detecting the presence of small intestinal bacterial overgrowth in a subject having at least one symptom associated with the diagnosis of one of these diseases.

ACTIVITY - Antiinflammatory; antidepressant; nootropic; tranquilizer; immunosuppressive; neuroprotective; dermatological.

Thirty subjects had previously received a diagnosis of chronic fatigue syndrome. Of these 30, 21 had small intestinal bacterial overgrowth (SIBO) as indicated by lactulose breath hydrogen testing (LBHT). Four out of the nine who did not have SIBO indicated had already received antibiotics. After treatment with neomycin (500 mg, twice daily for 10 days), 9 of the subjects with SIBO returned for LBHT and questionnaire. LBHT showed that all nine subjects experienced at least partial eradication of SIBO, and symptoms of bloating and fatigue were substantially improved.

MECHANISM OF ACTION - None given.

USE - The methods are used for the diagnosis and treatment of irritable bowel syndrome, **fibromyalgia**, chronic fatigue syndrome, depression, attention deficit/hyperactivity disorder, an autoimmune disease (e.g. multiple sclerosis or systemic lupus erythematosus) or Crohn's disease (claimed).

ADVANTAGE - The method diagnoses and treats the underlying causal factor of the diseases.
Dwg.0/2

L72 ANSWER 5 OF 6 HCAPLUS COPYRIGHT 2002 ACS DUPLICATE 2
1997:640529 Document No. 127:318776 Triarylmethane compounds for treatment of sickle cell disease. Brugnara, Carlo; Halperin, Jose; Bellot, Emile M., Jr.; Froimowitz, Mark; Lombardy, Richard J.; Clifford, John J.; Gao, Ying-duo; Haidar, Reem M.; Kelleher, Eugene W.; Kher, Falguni M.; Moussa, Adel M.; et al. (President and Fellows of Harvard College, USA; Children's Medical Center Corp.; Ion Pharmaceuticals, Inc.). PCT Int. Appl. WO 9734589 A1 19970925, 106 pp. DESIGNATED STATES: W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH, HU, IL, IS, JP, KG, KP, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, TJ, TM, TR, TT, UA, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 1997-US4551 19970319. PRIORITY: US 1996-618759 19960320; US 1996-618952 19960320; US 1996-618762 19960320; US 1996-618760 19960320.

AB The invention provides a class of chem. compds. useful in the treatment of both sickle cell disease and diseases characterized by unwanted or abnormal cell proliferation. The active compds. are substituted triarylmethane compds., or analogs where one or more aryl groups is replaced with a heteroaryl, cycloalkyl, or heterocycloalkyl group, and/or the tertiary C atom is replaced with a different atom such as Si, Ge, N, or P. The compds. inhibit mammalian cell proliferation, inhibit the Gardos channel of erythrocytes, reduce sickle erythrocyte dehydration, and/or delay the occurrence of erythrocyte sickling or deformation. Some of the compds. are novel and/or are prepd. in examples, while other compds. were obtained com. A total of 90 compds. were tested. For instance, reaction of 2-ClC₆H₄CPh₂Cl with Cu cyanide at 150.degree. in the absence of solvent gave 66% title nitrile 2-ClC₆H₄CPh₂CN (I). The IC₅₀ of I for inhibiting the Gardos channel of erythrocytes was 0.048 .mu.M (cf. 0.046 for clotrimazole), and that for inhibiting mitogen-induced cell proliferation in vitro was 2.20 .mu.M. Addnl. activity studies (animal and human) of clotrimazole and its triarylmethane metabolites are described.

L72 ANSWER 6 OF 6 MEDLINE DUPLICATE 3
84153862 Document Number: 84153862. PubMed ID: 6704108. Beta-lapachone greatly enhances MMS lethality to human **fibroblasts**. Boorstein R J; Pardee A B. BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS, (1984 Feb 14) 118 (3) 828-34. Journal code: 0372516. ISSN: 0006-291X. Pub. country: United States. Language: English.

AB beta-Lapachone is a naturally occurring **tricyclic** O-naphthoquinone. At microM concentrations it did not substantially affect viability, growth or DNA synthesis of cultured undamaged human **fibroblasts**. Cells exposed to minimally toxic concentrations of methyl **methane** sulfonate were strongly inhibited in these properties by beta-lapachone. The effects were not reversed by further incubation in the absence of beta-lapachone and were equal for initially quiescent or growing cells. Thus inhibitions were specific for damaged cells and did not involve replicative DNA synthesis. Inhibition of DNA strand break repair was demonstrated by alkaline elution, but unscheduled DNA synthesis was not inhibited. We propose that beta-lapachone inhibits a ligation step of DNA repair, in a manner perhaps similar to that reported for carbamoylating nitrosoureas. Other repair inhibitors differ significantly from beta-lapachone in their modes of action.

=> s l71 and ((triary? or tricycl?)(l)methane or cellular prolifer? or inflam?
disease or fibro? or lymphocy? or glomerulonephri?)

L73 0 FILE MEDLINE
L74 2 FILE BIOSIS
L75 5 FILE HCAPLUS
L76 0 FILE EMBASE
L77 0 FILE JICST-EPLUS
L78 8 FILE WPIDS

TOTAL FOR ALL FILES

L79 15 L71 AND ((TRIARY? OR TRICYCL?)(L) METHANE OR CELLULAR PROLIFER?
OR INFLAM? DISEASE OR FIBRO? OR LYMPHOCY? OR GLOMERULONEPHRI?)

=> s l79 not l64

L80 0 FILE MEDLINE
L81 1 FILE BIOSIS
L82 4 FILE HCAPLUS
L83 0 FILE EMBASE
L84 0 FILE JICST-EPLUS
L85 6 FILE WPIDS

TOTAL FOR ALL FILES

L86 11 L79 NOT L64

=> dup rem l86

PROCESSING COMPLETED FOR L86

L87 7 DUP REM L86 (4 DUPLICATES REMOVED)

=> d cbib abs 1-7

L87 ANSWER 1 OF 7 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE 1
2000:383154 Document No.: PREV200000383154. **Triaryl methane**

compounds and analogues thereof useful for the treatment or prevention of
sickle cell disease or diseases characterized by abnormal cell
proliferation. **Brugnara, Carlo (1)**; Halperin, Jose; Bellot,
Emile M.; Froimowitz, Mark; Lombardy, Richard John; Clifford, John J.;
Gao, Ying-Duo; Haidar, Reem M.; Kelleher, Eugene W.; Kher, Falguni M.;
Moussa, Adel M.; Sachdeva, Yesh P.; Sun, Minghu; Taft, Heather N.. (1)
Newton Highlands, MA USA. ASSIGNEE: Children's Medical Center Corporation;
Ion Pharmaceuticals, Inc., New York, NY, USA. Patent Info.: US 6028103
February 22, 2000. Official Gazette of the United States Patent and
Trademark Office Patents, (Feb. 22, 2000) Vol. 1231, No. 4, pp. No
pagination. e-file. ISSN: 0098-1133. Language: English.

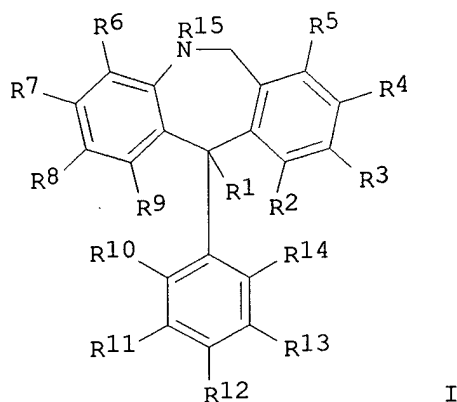
AB The present invention provides a class of chemical compounds useful as
efficacious drugs in the treatment of sickle cell disease and diseases
characterized by unwanted or abnormal cell proliferation. The active
compounds are substituted **triaryl methane** compounds or
analogues thereof where one or more of the aryl groups is replaced with a
heteroaryl, cycloalkyl or heterocycloalkyl group and/or the tertiary
carbon atom is replaced with a different atom such as Si, Ge, N or P. The
compounds inhibit mammalian cell proliferation, inhibit the Gardos channel
of erythrocytes, reduce sickle erythrocyte dehydration and/or delay the
occurrence of erythrocyte sickling or deformation.

L87 ANSWER 2 OF 7 HCAPLUS COPYRIGHT 2002 ACS DUPLICATE 2
1999:355731 Document No. 131:18936 Preparation of 11-phenyldibenzazepines
for the treatment sickle cell disease, **inflammatory**
diseases characterized by abnormal cell proliferation, diarrhea
and scour.. **Brugnara, Carlo**; Halperin, Jose; Bellot, Emile M.,

Searched by: Mary Hale 308-4258 CM-1 1E01

Jr.; Froimowitz, Mark; Lombardy, Richard John; Clifford, John J.; Gao, Ying-Duo; Haidar, Reem M.; Kelleher, Eugne W.; Kher, Falguni M.; Moussa, Adel M.; Sachdeva, Yesh P.; Sun, Minghua; Taft, Heather N.; Lencer, Wayne I.; Alper, Seth (Children's Medical Center Corporation, USA; President and Fellows of Harvard College; Ion Pharmaceuticals, Inc.). PCT Int. Appl. WO 9926628 A1 19990603, 92 pp. DESIGNATED STATES: W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG.
(English). CODEN: PIXXD2. APPLICATION: WO 1998-US24967 19981120.
PRIORITY: US 1997-975594 19971120; US 1998-159333 19980923; US 1998-159337 19980923.

GI



AB A method for inhibiting unwanted **cellular proliferation** assocd. with **inflammatory disease** comprises administration of title compds. [I; R1 = R', (substituted) aryl; R2, R3, R4 = R', OR', SR', halo, trihalomethyl; R3R4 = arylene; R5-R14 = R', halo, trihalomethyl; R15 = R'', COR'', CSR'', COCOOR'', etc.; R' = H, alkyl, alkenyl, alkynyl; R'' = R', (substituted) aryl, aralkyl, alkylaryl]. I are specific, potent and safe inhibitors of the Ca²⁺-activated potassium channel (Gardos channel) of erythrocytes, of mammalian cell proliferation and/or of secretagogue-stimulated transepithelial electrogenic chloride secretion in intestinal cells. I can be used to reduce sickle erythrocyte dehydration and/or delay the occurrence of erythrocyte sickling or deformation in situ as a therapeutic approach towards the treatment or prevention of sickle cell disease. I can also be used to inhibit mammalian cell proliferation in situ as a therapeutic approach towards the treatment or prevention of diseases characterized by abnormal cell proliferation. I can also be used to inhibit chloride secretion in intestinal cells as a therapeutic approach towards the treatment of diarrhea and scours. Thus, I (R1-R15 = H), K₂CO₃, and MeO₂CCl were refluxed 12 h in MeCN to give 48% I (R1-R14 = H; R15 = MeO₂C). The latter inhibited the Gardos channel with IC₅₀ = 0.0850-0.093 .mu.M.

L87 ANSWER 3 OF 7 WPIDS (C) 2002 THOMSON DERWENT
AN 1999-347682 [29] WPIDS

Searched by: Mary Hale 308-4258 CM-1 1E01

AB WO 9926929 A UPAB: 19990723

NOVELTY - Substituted 11-phenyl-dibenzazepine derivatives (I) are new.

DETAILED DESCRIPTION - Substituted 11-phenyl-dibenzazepine derivatives of formula (I) and their salts and hydrates are new:

R1 = R' or 6-20C aryl optionally substituted with Q;

R2, R3, R4 = R', -OR', -SR', halo or trihalomethyl; or R3+R4 form 6-20C aryleneo;

R5-R14 = R', halo or trihalomethyl;

R15 = R, -C(O)R, -C(S)R, -C(O)OR, -C(S)OR, -C(O)SR, -C(S)SR, C(O)N(R')2, -C(S)N(R)2, -C(O)C(O)R, -C(S)C(O)R, C(O)C(S)R, -C(S)C(S)R, -C(O)C(O)OR, -C(S)C(O)OR, C(O)C(S)OR, C(O)C(O)SR, -C(S)C(S)OR, -C(S)C(O)SR, C(O)C(S)SR, C(S)C(S)SR, -C(O)C(O)N(R)2, -C(S)C(O)N(R)2, C(O)C(S)N(R)2 or -C(S)C(S)N(R)2;

R' = H, 1-6C alkyl, 2-6C alkenyl or 2-6C alkynyl;

R = R', 6-20C aryl optionally substituted with Q, or 6-26C alkaryl optionally substituted with Q;

Q = CN, -OR', -SR', NO2, -NR'2, halo, 1-6C alkyl, 2-6C alkenyl, 2-6C alkynyl or trihalomethyl; provided that when R1 and R15 are both H, at least 1 of R1-R14 is other than H, R8 is other than H or Cl and at least 1 of R2-R5 is other than -OMe.

ACTIVITY - Antiproliferative; antineoplastic. Antiproliferative assays were carried out using cancer cell lines. Results for N-(4'-nitrobenzoyl)-11-(2'-chlorophenyl)-5,6 dihydro-11H-dibenz(b,e)azepine (Ia) and anti-cancer agent VP-16 respectively after incubation for 3 days were: against A549, 1.8 and 2.3 μ M; HT29, less than 1.25 and 20 μ M; and MCF7, 2.1 and less than 2.5 μ M.

MECHANISM OF ACTION - Inhibitors of Gardos channel (Ca²⁺ activated potassium channel) of erythrocytes; inhibitors of mitogen-induced cell proliferation. Compounds including N-methyl-11-phenyl-5,6-dihydro-11H-dibenz(b,e)azepine (Ib) and clotrimazole were tested in-vitro for (a) % inhibition of the Gardos channel (10 μ M compound) and IC50 as described in Brugnara et al., 1993, J.Biol.Chem. 268(12):8760-8768; and (b) % inhibition of mitogen-induced cell proliferation (10 μ M) and IC50 as described in Benzaquen et al., 1995, Nature Medicine 1:534-540. Results were (a) for (Ib) IC50 1.30 μ M and 99% inhibition compared with IC50 0.046 μ M and 99.3% inhibition for clotrimazole; and (b) for (Ib) IC50 5.2 μ M and 99% inhibition, compared with IC50 0.626 μ M and 93% inhibition for clotrimazole.

USE - (I) are used for treating or preventing disorders characterized by abnormal cell proliferation (particularly in endothelial, **fibrotic** or vascular smooth muscle cells), e.g. cancer; a blood vessel proliferative disorder; a **fibrotic** disorder; an arteriosclerotic condition; or a dermatological disease such as keloids, hypertonic sacars, seborrheic dermatosis, papilloma virus infection, eczema or actinic keratosis; or Kaposi's sarcoma (all claimed). (I) are also useful for treating or preventing sickle cell disease. (I) can be administered with other active agents.

ADVANTAGE - (I) have reduced toxicity compared with clotrimazole and other antimycotic agents.
Dwg.0/0

L87 ANSWER 4 OF 7 WPIDS (C) 2002 THOMSON DERWENT

AN 1999-347610 [29] WPIDS

AB WO 9926624 A UPAB: 19990723

NOVELTY - Substituted diphenyl indanone, indane and indole compounds can be used to treat **inflammatory diseases** characterized by abnormal cell proliferation and inhibit chloride secretion in the treatment of diarrhea and scours.

DETAILED DESCRIPTION - The use is claimed of substituted diphenyl indanone, indane and indole compounds of formula (I) for treating

inflammatory diseases characterized by abnormal cell proliferation (but not cancer, actinic keratosis or Kaposi's sarcoma) and treating diarrhea and scours.

m = 0-4;

n = 0-5;

dotted line = single or double bond;

X = C or N;

Y = absent, 1-6C alkyl, 2-6C alkenyl or 2-6C alkynyl;

R1 = absent, OR, SR, O, S, N OR, OC(O)R, SC(O)R, OC(O)R or SC(O)R, or R1+R2 form a 3-8 membered heterocycloalkyl optionally substituted with Q;

R2, R3 = absent or H;

R4 = H, OR', SR', NR'2, CN, NO2, 3-8C cycloalkyl, 3-8 membered heterocycloalkyl, C(O)R', C(O)R', C(O)OR', C(O)OR', C(O)SR', C(S)SR', C(O)NR'2 or C(S)NR'2;

R5-R7 = halo, R', OR', SR', NR'2, ONR'2, SNR'2, NO2, CN, C(O)R', C(S)R', C(O)OR', C(O)SR', C(S)OR', CS(S)R', C(O)NR'2, C(S)NR'2, C(O)NR'(OR'), C(S)NR'(OR'), C(O)NR'(SR'), C(S)NR'(SR'), CH(CN)2, CH(C(O)R')2, CH(C(S)R')2, CH(C(O)OR')2, CH(C(S)OR')2, CH(C(O)SR')2 or CH(C(S)SR')2;

R = H, 1-6C alkyl, 2-6C alkenyl, 2-6C alkynyl, 5-20C aryl optionally substituted with Q', or 6-26C alkaryl optionally substituted with Q';

Q = CN, NO2, NR'2, OR', C(O)NR'2, C(S)NR'2, C(O)OR', C(S)OR', C(O)SR', C(S)SR' or trihalomethyl;

Q' = halo, C(O)R', C(S)R', C(O)OR', C(S)OR', C(O)SR', C(S)SR', C(O)NR'2, C(S)NR'2 or trihalomethyl; and

R' = H, 1-6C alkyl, 2-6C alkenyl or 2-6C alkynyl.

INDEPENDENT CLAIMS are included for the following:

(a) the use of an optionally substituted 3,3-diphenyl indanone, an optionally substituted indane or an optionally substituted (3H)-indole derivative, and analogues of these compounds where the atoms at ring positions 1 and 2 are connected via a double bond, for inhibiting chloride secretion and treating diarrhea; and for treating scours;

(b) a veterinary preparation comprising a compound described in (a) and an anti-scours agent; and

(c) a composition for treating diarrhea comprising a compound described in (a) and an anti-diarrheal agent.

ACTIVITY - Antiproliferative.

MECHANISM OF ACTION - Inhibitors of Gardos channel (Ca²⁺ activated potassium channel) of erythrocytes; inhibitors of mammalian cell proliferation; inhibitors of chloride secretion in intestinal cells.

1-Hydroxy-3,3-diphenylindane (10 micro M) inhibited Gardos channel activity by 100%, with an IC₅₀ value of 0.819 micro M (corresponding values for clotrimazole are 99.3% and 0.046 micro M).

USE - (I) can be used to treat **inflammatory diseases** including proliferative **glomerulonephritis**; lupus erythematosus; scleroderma; temporal arteritis; thromboangiitis obliterans; mucocutaneous lymph node syndrome; asthma; graft versus host; inflammatory bowel disease; multiple sclerosis; rheumatoid arthritis; thyroiditis; Grave's disease; antigen-induced airway hyperactivity; pulmonary eosinophilia; Guillain-Barre syndrome; allergic rhinitis; myasthenia gravis; human T-lymphotrophic virus type I-associated myelopathy; herpes simplex encephalitis; inflammatory myelopathies; atherosclerosis; and Goodpasture's syndrome; also diarrhea and scours (in e.g. horse, cow, pig or goat).

Dwg.0/4

L87 ANSWER 5 OF 7 WPIDS (C) 2002 THOMSON DERWENT

AN 1999-404733 [34] WPIDS

AB WO 9926611 A UPAB: 20020502

Searched by: Mary Hale 308-4258 CM-1 1E01

NOVELTY - Substituted 3,3-diphenyl indanone, indane and indole derivatives (I) are new.

DETAILED DESCRIPTION - Indanone, indane and indole derivatives of formula (I) and their salts and hydrates are new.

rings a, b and c are optionally substituted by 1-5 R5 groups, 1-5 R6 groups and 1-4 R7 groups respectively;

X = C or N;

Y = direct bond, 1-6C alkylene, 2-6C alkenylene or 2-6C alkynylene;

R1 = -OR, -SR, =O, =S, =NOR, O- C(O)R, -S-C(O)R, -O-C(O)R or -S-C(O)R, or is absent;

R2, R3 = H or are absent;

or R1+R2 = group completing 3-8 membered heterocycloalkyl (optionally substituted by Q);

R4 = H, -OR', -SR', -NR'2, CN, NO2, 3-8C cycloalkyl, 3-8 membered heterocycloalkyl, -C(O)R', -C(O)R', C(O)OR', -C(O)OR', -C(O)SR', -C(S)SR', -C(O)NR'2 or -C(S)NR'2;

R5 - R7 = halo, R', -OR', -SR', -NR'2, -ONR'2, -SNR'2, NO2, CN, -C(O)R', -C(S)R', -C(O)OR', -C(O)SR', -C(S)OR', CS(S)R', -C(O)NR'2, -C(S)NR'2, -C(O)NR'(OR'), -C(S)NR'(OR'), C(O)NR'(SR'), -C(S)NR'(SR'), -CH(CN)2, -CH(C(O)R')2, CH(C(S)R')2, -CH(C(O)OR')2, -CH(C(S)OR')2, -CH(C(O)SR')2 or CH(C(S)SR')2;

R = H, 1-6C alkyl, 2-6C alkenyl or 2-6C alkynyl; or 5-20C aryl or 6-26C alkaryl (optionally substituted by Q');

Q = CN, NO2, -NR'2, -OR', C(O)NR'2, -C(S)NR'2, -C(O)OR', -C(S)OR', -C(O)SR', -C(S)SR' or trihalomethyl;

Q' = halo, -C(O)R', -C(S)R', -C(O)OR', -C(S)OR', -C(O)SR', -C(S)SR', -C(O)NR'2, -C(S)NR'2 or trihalomethyl;

R' = H, 1-6C alkyl, 2-6C alkenyl or 2-6C alkynyl;

broken line = single or double bond;

provided that

(1) if X = C and R1 = =O or OH, then at least one of R5- R7 is other than H, or Y is present or R4 is other than H; and

(2) if X = N, broken line = double bond, R1- R3 and Y = bond, then R4 is other than NH2.

ACTIVITY - Antiproliferative; antineoplastic; dermatological; cardiovascular. Antiproliferative assays were carried out using cancer cell lines. Results for 1-N-oxime-3,3-diphenylindane (Ia) and anticancer agent VP-16 respectively after incubation for 3 days were: against A549, 8.5 and 2.3 mu M; HT29, less than 2.5 and 20 mu M; and MCF7, less than 2.5 mu M for both.

MECHANISM OF ACTION - Erythrocyte Gardos channel (calcium ion activated potassium channel) inhibitor; mitogen-induced cell proliferation inhibitor. Compounds including 1-N-oxime-3,3-diphenylindane (Ia) and clotrimazole were tested in vitro for (a) % inhibition of the Gardos channel (10 mu M compound) and IC50 as described in J. Biol. Chem. 268 (12), 8760-8768, 1993; and (b) % inhibition of mitogen-induced cell proliferation (10 mu M) and IC50 as described in Nature Medicine 1: 534-540, 1995.

Results were (a) for (Ia) IC50 1.35 mu M and 100% inhibition compared with IC50 0.046 mu M and 99.3% inhibition for clotrimazole; and (b) for (Ia) IC50 2.6 mu M and 99% inhibition, compared with IC50 0.626 mu M and 93% inhibition for clotrimazole.

USE - For treating or preventing disorders characterized by abnormal cell proliferation (particularly in endothelial, **fibrotic** or vascular smooth muscle cells), specifically cancer, blood vessel proliferative disorders, **fibrotic** disorders, arteriosclerotic conditions, dermatological diseases (especially keloids, hypertonic scars, seborrheic dermatosis, papilloma virus infection, eczema or actinic keratosis) or Kaposi's sarcoma (all claimed). (I) are also useful for treating or preventing sickle cell disease.

ADVANTAGE - (I) are specific, potent and safe inhibitors of the calcium ion activated potassium channel (Gardos channel) of erythrocytes (particularly sickle erythrocytes) and/or of mammalian cell proliferation. They have reduced toxicity compared with clotrimazole and similar agents. Dwg.0/0

L87 ANSWER 6 OF 7 HCAPLUS COPYRIGHT 2002 ACS DUPLICATE 3
1994:622006 Document No. 121:222006 Imidazoles for treatment of arteriosclerosis. Halperin, Jose; **Brugnara, Carlo** (President and Fellows of Harvard College, USA). PCT Int. Appl. WO 9418968 A1 19940901, 26 pp. DESIGNATED STATES: W: AU, CA, JP; RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE. (English). CODEN: PIXXD2. APPLICATION: WO 1994-US1749 19940218. PRIORITY: US 1993-18835 19930218.

AB An imidazole which inhibits the proliferation of endothelial cells, vascular smooth muscle cells and **fibroblasts** and inhibits the Ca++-activated K channel, is used for the treatment of arteriosclerotic conditions. For example, dose-response inhibitions of DNA synthesis by clotrimazole was tested using rat vascular smooth muscle cells.

L87 ANSWER 7 OF 7 HCAPLUS COPYRIGHT 2002 ACS
1986:455276 Document No. 105:55276 Bradykinin and vasopressin stimulate sodium-potassium-chloride cotransport in cultured endothelial cells. Brock, Tommy A.; **Brugnara, Carlo**; Canessa, Mitzy; Gimbrone, Michael A., Jr. (Dep. Pathol., Brigham Women's Hosp., Boston, MA, 02115, USA). American Journal of Physiology, 250(6, Pt. 1), C888-C895 (English) 1986. CODEN: AJPHAP. ISSN: 0002-9513.

AB A Na+-K+-Cl- cotransporter was characterized in vascular endothelial cells (EC) cultured from different blood vessels and species, that is inhibited by the diuretics furosemide and bumetanide (50% inhibitory concn. for 86Rb+ influx .apprx.20 .mu.M and 0.5 .mu.M, resp.). Inward 86Rb+ influx mediated via this pathway is greater than 86Rb+ influx transported by the Na+-K+ pump in cultured EC from bovine and pig aorta, bovine vena cava, and baboon cephalic vein but not in human umbilical or saphenous vein EC. External Na+- or Cl--stimulated, ouabain-insensitive 86Rb+ influx is equal to furosemide- or bumetanide-sensitive 86Rb+ influx. Ouabain-insensitive 22Na+ influx is also partially inhibited by these drugs and stimulated by increasing external K+ or Cl-. Net Na+ extrusion occurs via the Na+-K+-Cl- cotransporter in the absence of external K+, whereas net Na+ influx occurs at higher external K+ (>1 mM). Maximal concns. (100 nM) of bradykinin [58-82-2] and vasopressin [11000-17-2] increase the initial rate of bumetanide-sensitive 86Rb+ influx by .apprx.60% and 70% (50% effective concn. .apprx.1 nM and 0.6 nM, resp.). Addn. of either EGTA or LaCl3 (to block Ca2+ influx) prevents bradykinin-stimulated 86Rb+ influx. When intracellular Ca2+ is elevated by using ionomycin (100 nM), a Ca2+ ionophore, bumetanide-sensitive 86Rb+ influx increases .apprx.2-fold. In contrast, isoproterenol [60-92-4] (100 .mu.M) and forskolin (50 .mu.M), adenylate cyclase [9012-42-4] stimulators, decrease furosemide-sensitive 86Rb+ influx. Thus, in certain types of cultured EC, a Na+-K+-Cl- cotransporter mediates a fraction of K+ influx quant. as important as the Na+-K+ pump (ouabain-sensitive 86Rb+ influx) and appears to be modulated by Ca2+ and cyclic nucleotides.

=> log y

COST IN U.S. DOLLARS
FULL ESTIMATED COST

| SINCE FILE | TOTAL |
|------------|---------|
| ENTRY | SESSION |
| 33.59 | 962.82 |

Searched by: Mary Hale 308-4258 CM-1 1E01

| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE | TOTAL |
|--|------------|---------|
| | ENTRY | SESSION |
| CA SUBSCRIBER PRICE | -2.48 | -13.10 |

STN INTERNATIONAL LOGOFF AT 13:58:38 ON 08 NOV 2002

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

REFERENCE 2: 132:166005 Triaryl methane compounds and analogues thereof useful for the treatment or prevention of sickle cell disease or diseases characterized by abnormal cell proliferation. Brugnara, Carlo; Halperin, Jose; Bellot, Emile M., Jr.; Froimowitz, Mark; Lombardy, Richard John; Clifford, John J.; Gao, Ying-duo; Haidar, Reem M.; Kelleher, Eugene W.; Kher, Falguni M.; Moussa, Adel M.; Sachdeva, Yesh P.; Sun, Minghua; Taft, Heather N. (Children's Medical Center Corporation, USA; Ion Pharmaceuticals, Inc.). U.S. US 6028103 A 20000222, 95 pp., Cont.-in-part of U.S. Ser. No. 618,952. (English). CODEN: USXXAM. APPLICATION: US 1997-822550 19970319. PRIORITY: US 1996-618952 19960320; US 1996-618760 19960320.

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The present invention provides a class of chem. compds. such as I [n = 0-4; R1 = H, CN, OR, etc.; R2 = F, Cl, Br, I; R3 = R, OR, SR; R4 = H, NR2; R5 = H, F, Cl, Br, I; R = H, alkyl, alkenyl, etc.], II [n = 0-4; R1 = NR2, COR, CSR, etc.; R2-R4 = F, Cl, Br, I; R = H, alkyl, alkenyl, etc.] and III [n = 0-4; Ar1 = Ph, cyclohexyl; R1 = NR2, CSNR2, CONR2, etc.; R = H, alkyl, alkenyl, etc.] which inhibit mammalian cell proliferation, inhibit the Gardos channel of erythrocytes, reduce sickle erythrocyte dehydration and/or delay the occurrence of erythrocyte sickling or deformation, and therefore are useful as efficacious drugs in the treatment of sickle cell disease and diseases characterized by unwanted or abnormal cell proliferation. The active compds. are substituted triaryl methane compds. or analogs thereof where one or more of the aryl groups is replaced with a heteroaryl, cycloalkyl or heterocycloalkyl group and/or the tertiary carbon atom is replaced with a different atom such as Si, Ge, N or P. Prepn. of some of compds. I-III was presented. Biol. data (e.g., inhibition of mitogen-induced cell proliferation and inhibition of Gardos channel) for all exemplified compds. I-III were given.

REFERENCE 3: 127:318776 Triarylmethane compounds for treatment of sickle cell disease. Brugnara, Carlo; Halperin, Jose; Bellot, Emile M., Jr.; Froimowitz, Mark; Lombardy, Richard J.; Clifford, John J.; Gao, Ying-duo;

Haidar, Reem M.; Kelleher, Eugene W.; Kher, Falguni M.; Moussa, Adel M.; et al. (President and Fellows of Harvard College, USA; Children's Medical Center Corp.; Ion Pharmaceuticals, Inc.). PCT Int. Appl. WO 9734589 A1 19970925, 106 pp. DESIGNATED STATES: W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH, HU, IL, IS, JP, KG, KP, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, TJ, TM, TR, TT, UA, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 1997-US4551 19970319. PRIORITY: US 1996-618759 19960320; US 1996-618952 19960320; US 1996-618762 19960320; US 1996-618760 19960320.

AB The invention provides a class of chem. compds. useful in the treatment of both sickle cell disease and diseases characterized by unwanted or abnormal cell proliferation. The active compds. are substituted triarylmethane compds., or analogs where one or more aryl groups is replaced with a heteroaryl, cycloalkyl, or heterocycloalkyl group, and/or the tertiary C atom is replaced with a different atom such as Si, Ge, N, or P. The compds. inhibit mammalian cell proliferation, inhibit the Gardos channel of erythrocytes, reduce sickle erythrocyte dehydration, and/or delay the occurrence of erythrocyte sickling or deformation. Some of the compds. are novel and/or are prepd. in examples, while other compds. were obtained com. A total of 90 compds. were tested. For instance, reaction of 2-ClC6H4CPh2Cl with Cu cyanide at 150.degree. in the absence of solvent gave 66% title nitrile 2-ClC6H4CPh2CN (I). The IC50 of I for inhibiting the Gardos channel of erythrocytes was 0.048 .mu.M (cf. 0.046 for clotrimazole), and that for inhibiting mitogen-induced cell proliferation in vitro was 2.20 .mu.M. Addnl. activity studies (animal and human) of clotrimazole and its triarylmethane metabolites are described.

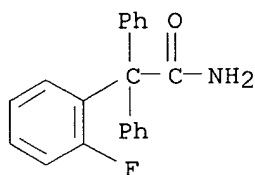
L49 ANSWER 10 OF 10 REGISTRY COPYRIGHT 2002 ACS

RN 82757-34-4 REGISTRY

CN Benzeneacetamide, 2-fluoro-.alpha.,.alpha.-diphenyl- (9CI) (CA INDEX NAME)

MF C20 H16 F N O

LC STN Files: CA, CAPLUS, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2 REFERENCES IN FILE CA (1962 TO DATE)

2 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU,

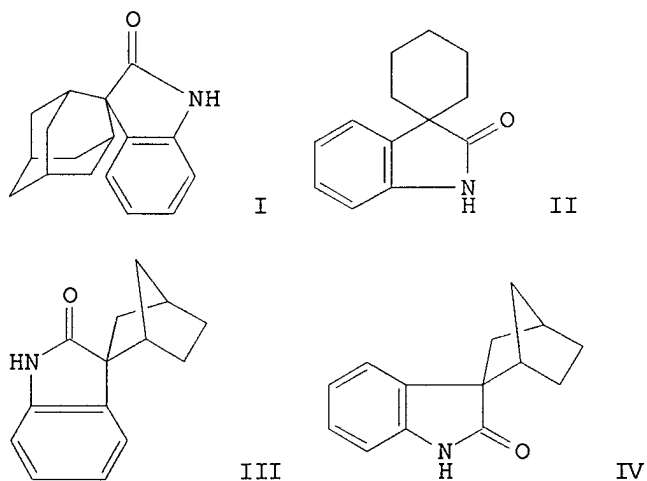
Searched by: Mary Hale 308-4258 CM-1 1E01

ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

REFERENCE 2: 97:109824 Two new stereochemically complementary oxindole syntheses. Fleming, Ian; Loreto, Maria Antonietta; Michael, Joseph P.; Wallace, Ian H. M. (Chem. Lab., Univ. Cambridge, Cambridge, CB2 1EW, UK). Tetrahedron Lett., 23(19), 2053-6 (English) 1982. CODEN: TELEAY. ISSN: 0040-4039.

GI



AB Two routes are reported for the conversion of ketones to oxindoles. The 1st route involved sequential addn. reaction of the ketone with 2-LiC6H4NLiCHO in THF at -105.degree., followed by sequential cyanation, cyclization, and hydrolysis to give the oxindole. The route 2 comprised sequential Wittig reaction of the ketone with 2-FC6H4CH:PPh3, epoxidn., rearrangement, amidation, and intramol. cyclocondensation reaction. E.g., using route 1, the oxindole I was prepd. in 79% from adamantanone, and using route 2 the oxindole II was prepd. in moderate yield from cyclohexanone. From norbornanone, route 1 gave the Wallace oxindole III and route 2 gave the Loreto oxindole IV.

=> fil caol;s 149

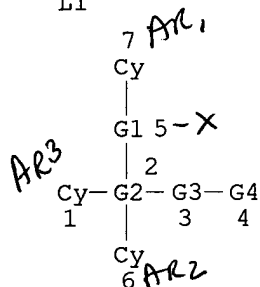
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FULL ESTIMATED COST

| | |
|------------|---------|
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| ENTRY | SESSION |
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Searched by: Mary Hale 308-4258 CM-1 1E01

Sackey
942258

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can;d 111 que stat
L1 STR



x= REP G1=(1-3) C
y= VAR G2=C/N/P/SI/GE
n= REP G3=(0-4) CH2
R1= VAR G4=X/H/O/S/N/C
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
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STEREO ATTRIBUTES: NONE
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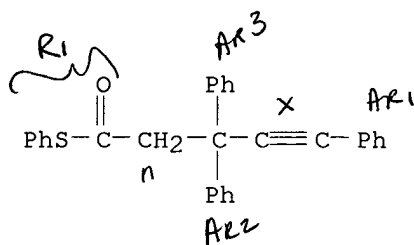
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BATCH **INCOMPLETE**
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PROJECTED ANSWERS: EXCEEDS 35714

2 ANSWERS
*not searchable
using above
query*

sample examples:

L26 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2002 ACS
RN 459126-26-2 REGISTRY
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ester (9CI) (CA INDEX NAME)
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SR CA
LC STN Files: CA, CAPLUS

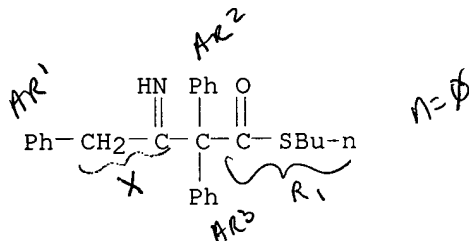


PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 137:232420

L26 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2002 ACS
RN 32188-89-9 REGISTRY
CN Butyric acid, 3-imino-2,2,4-triphenylthio-, S-butyl ester (8CI) (CA INDEX NAME)
MF C26 H27 N O S
LC STN Files: BEILSTEIN*, CA, CAPLUS
(*File contains numerically searchable property data)

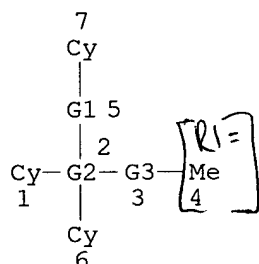


PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 75:5440

L3 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 7

STEREO ATTRIBUTES: NONE
L4 0 SEA FILE=REGISTRY SSS SAM L3

0.1% PROCESSED 1000 ITERATIONS

0 ANSWERS

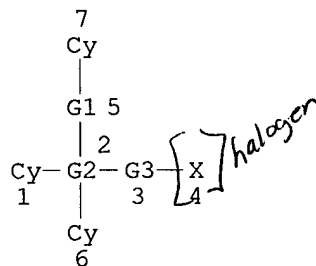
Searched by: Mary Hale 308-4258 CM-1 1E01

INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

*not searchable
too many iterations*

L5 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 7

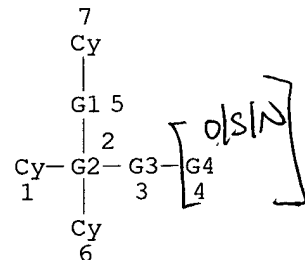
STEREO ATTRIBUTES: NONE
L6 0 SEA FILE=REGISTRY SSS SAM L5

0.1% PROCESSED 1000 ITERATIONS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

L8 STR



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 VAR G4=O/S/N
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 7

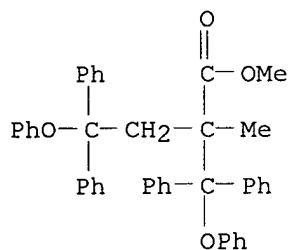
STEREO ATTRIBUTES: NONE
 L9 1 SEA FILE=REGISTRY SSS SAM L8

0.1% PROCESSED 1000 ITERATIONS 1 ANSWERS
 INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
 SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
 BATCH **INCOMPLETE**
 PROJECTED ITERATIONS: EXCEEDS 1000000
 PROJECTED ANSWERS: EXCEEDS 17312

Sample examples:

L9 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS
 RN 118408-00-7 REGISTRY
 CN Benzenebutanoic acid, .alpha.-methyl-.gamma.-phenoxy-.alpha.-
 (phenoxydiphenylmethyl)-.gamma.-phenyl-, methyl ester (9CI) (CA INDEX
 NAME)
 MF C43 H38 O4
 SR CA
 LC STN Files: CA, CAPLUS



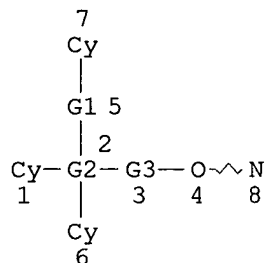
PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
 1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 110:58163

Searched by: Mary Hale 308-4258 CM-1 1E01

L10 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

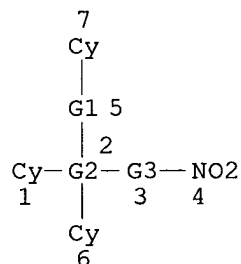
GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 8

STEREO ATTRIBUTES: NONE
L11 0 SEA FILE=REGISTRY SSS SAM L10

1.2% PROCESSED 1000 ITERATIONS 0 ANSWERS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

=> d 113 que stat;d 113 que stat;d 115 que stat;d ide can 115;d 117 que stat;d 119
que stat;d 121 que stat
L12 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED

Searched by: Mary Hale 308-4258 CM-1 1E01

NUMBER OF NODES IS 7

STEREO ATTRIBUTES: NONE

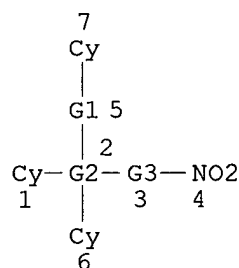
L13 0 SEA FILE=REGISTRY SSS SAM L12

1.7% PROCESSED 1000 ITERATIONS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

L12 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 7

STEREO ATTRIBUTES: NONE

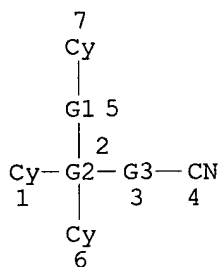
L13 0 SEA FILE=REGISTRY SSS SAM L12

1.7% PROCESSED 1000 ITERATIONS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

L14 STR



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 7

STEREO ATTRIBUTES: NONE
 L15 1 SEA FILE=REGISTRY SSS SAM L14

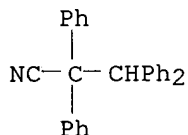
2.2% PROCESSED 1000 ITERATIONS
 INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
 SEARCH TIME: 00.00.01

1 ANSWERS

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
 BATCH **INCOMPLETE**
 PROJECTED ITERATIONS: 905497 TO 930943
 PROJECTED ANSWERS: 512 TO 1324

sample example

L15 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS
 RN 1249-27-0 REGISTRY
 CN Benzenepropanenitrile, .alpha.,.alpha.,.beta.-triphenyl- (9CI) (CA INDEX NAME)
 OTHER CA INDEX NAMES:
 CN Propionitrile, 2,2,3,3-tetraphenyl- (7CI, 8CI)
 MF C27 H21 N
 LC STN Files: BEILSTEIN*, CA, CAOLD, CAPLUS
 (*File contains numerically searchable property data)



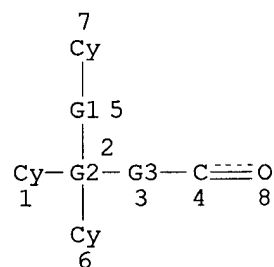
PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

Searched by: Mary Hale 308-4258 CM-1 1E01

6 REFERENCES IN FILE CA (1962 TO DATE)
6 REFERENCES IN FILE CAPLUS (1962 TO DATE)
2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 98:160451
REFERENCE 2: 87:5287
REFERENCE 3: 64:93117
REFERENCE 4: 64:93116
REFERENCE 5: 62:66318
REFERENCE 6: 62:66317

L16 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 8

STEREO ATTRIBUTES: NONE
L17 0 SEA FILE=REGISTRY SSS SAM L16

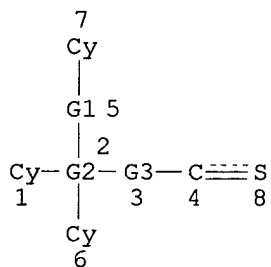
0.1% PROCESSED 1000 ITERATIONS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

L18 STR

Searched by: Mary Hale 308-4258 CM-1 1E01



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 8

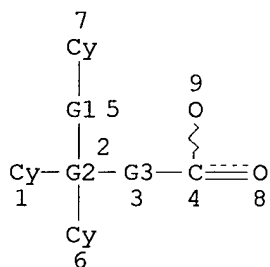
STEREO ATTRIBUTES: NONE
 L19 0 SEA FILE=REGISTRY SSS SAM L18

0.6% PROCESSED 1000 ITERATIONS
 INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
 SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
 BATCH **INCOMPLETE**
 PROJECTED ITERATIONS: EXCEEDS 1000000
 PROJECTED ANSWERS: EXCEEDS 0

L20 STR



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 9

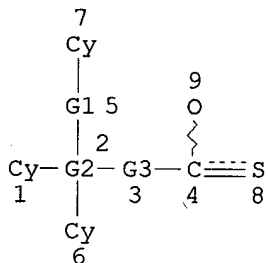
Searched by: Mary Hale 308-4258 CM-1 1E01

STEREO ATTRIBUTES: NONE
L21 0 SEA FILE=REGISTRY SSS SAM L20

0.3% PROCESSED 1000 ITERATIONS 0 ANSWERS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

=> d 123 que stat;d 126 que stat;d 1-2 ide cbib abs
L22 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 9

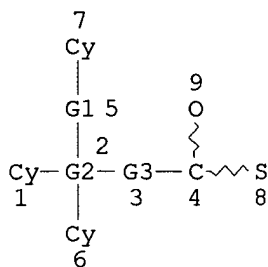
STEREO ATTRIBUTES: NONE
L23 0 SEA FILE=REGISTRY SSS SAM L22

15.6% PROCESSED 1000 ITERATIONS 0 ANSWERS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 123580 TO 133180
PROJECTED ANSWERS: 0 TO 0

*continue
further for
complete search
see L26 results*

L24 STR



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

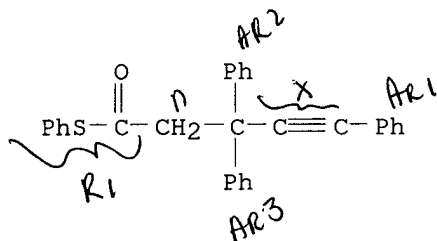
GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 9

STEREO ATTRIBUTES: NONE
 L26 2 SEA FILE=REGISTRY SSS FUL L24

100.0% PROCESSED 127186 ITERATIONS
 SEARCH TIME: 00.00.10

2 ANSWERS

L26 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2002 ACS
 RN 459126-26-2 REGISTRY
 CN Benzenepropanethioic acid, .beta.-phenyl-.beta.-(phenylethynyl)-, S-phenyl
 ester (9CI) (CA INDEX NAME)
 FS 3D CONCORD
 MF C29 H22 O S
 SR CA
 LC STN Files: CA, CAPLUS



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
 1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 137:232420 Iridium-Catalyzed Substitution of Propargylic-type
 Esters with Enoxysilanes. Matsuda, Isamu; Komori, Kenichi; Itoh, Kenji
 (Graduate School of Engineering, Department of Molecular Design and
 Engineering, Nagoya University, Chikusa, Nagoya, 464-8603, Japan).

Searched by: Mary Hale 308-4258 CM-1 1E01

Journal of the American Chemical Society, 124(31), 9072-9073 (English)
2002. CODEN: JACSAT. ISSN: 0002-7863. Publisher: American Chemical Society.

AB Propargylic-type acetates react readily with enoxysilanes in the presence of 1 mol % of [Ir(cod){P(OPh)₃}₂]OTf activated preliminarily with mol. H₂ to give .beta.-alkynyl ketones in high to excellent yields. Substitution at the propargyl carbon proceeds exclusively or selectively in most types of propargylic esters. Alternatively, the formation of the allenyl products is predominant in the reaction of esters which have two Ph groups on the propargyl carbon.

L26 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2002 ACS

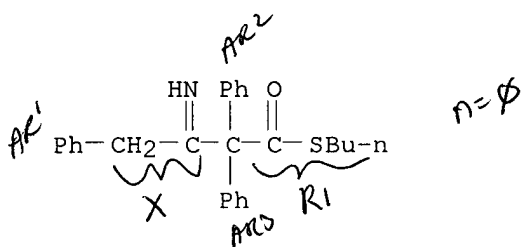
RN 32188-89-9 REGISTRY

CN Butyric acid, 3-imino-2,2,4-triphenylthio-, S-butyl ester (8CI) (CA INDEX NAME)

MF C26 H27 N O S

LC STN Files: BEILSTEIN*, CA, CAPLUS

(*File contains numerically searchable property data)



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)

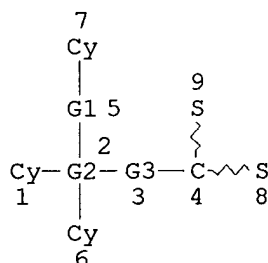
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 75:5440 Reactions of thioboronite. Mukaiyama, Teruaki; Inomata, Katsuhiko; Yamamoto, Shoji (Lab. Org. Chem., Tokyo Inst. Technol., Tokyo, Japan). Tetrahedron Lett. (16), 1097-100 (English) 1971. CODEN: TELEAY.

AB (EtS)₃B reacted with PhNCO to give the 1:2 adduct N-(ethylthiocarbonyl)-N,N'-diphenylurea and similar 1:2 adducts were obtained using Bu₂BSBu and PhNCO, cyclohexyl isocyanate, or EtNCO. PhNCS and dicyclohexylcarbodiimide gave 1:1 adducts, while reaction with PhCH₂CN gave a nitrile-thioboronite coordination complex, which gave acetamidine derivs. on treatment with isocyanates.

=> d l34 que stat;d ide cbib abs l34

L27 STR



REP G1=(1-3) C

VAR G2=C/N/P/SI/GE

Searched by: Mary Hale 308-4258 CM-1 1E01

REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

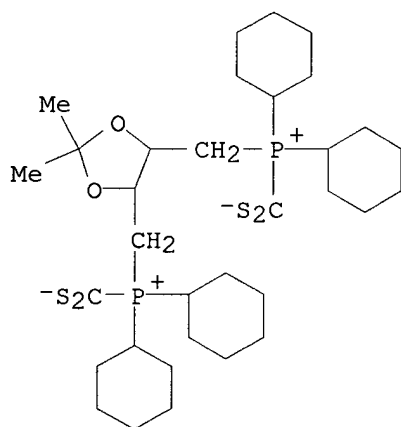
GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 9

STEREO ATTRIBUTES: NONE
L34 1 SEA FILE=REGISTRY SSS FUL L27

100.0% PROCESSED 85332 ITERATIONS
SEARCH TIME: 00.00.07

1 ANSWERS

L34 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS
RN 82239-69-8 REGISTRY
CN Phosphonium, [(2,2-dimethyl-1,3-dioxolane-4,5-
diyl)bis(methylene)]bis[dicyclohexyl(dithiocarboxy)-, bis(inner salt),
(4R-trans)- (9CI) (CA INDEX NAME)
MF C33 H56 O2 P2 S4
LC STN Files: CA, CAPLUS, CASREACT

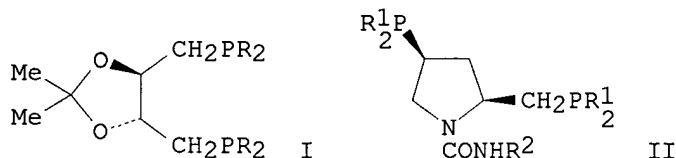


2 REFERENCES IN FILE CA (1962 TO DATE)
2 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 112:158338 Preparation of optically active peralkyldiphosphines and their use, as the rhodium(I) complex, in the asymmetric catalytic hydrogenation of ketones. Tani, Kazuhide; Suwa, Kenichi; Tanigawa, Eiichi; Ise, Tomokazu; Yamagata, Tsuneaki; Tatsuno, Yoshitaka; Otsuka, Sei (Fac. Eng. Sci., Osaka Univ., Osaka, 560, Japan). Journal of Organometallic Chemistry, 370(1-3), 203-21 (English) 1989. CODEN: JORCAI. ISSN: 0022-328X.

GI

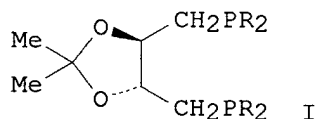
Searched by: Mary Hale 308-4258 CM-1 1E01



AB Two types of the optically active peralkyldiphosphine, 2,3-O-isopropylidene-2,3-dihydroxy-1,4-bis(dialkylphosphino)butane (Rdiop) I [R = Et, Me₂CH, cyclohexyl (Cy)] and N-(N'-substituted carbamoyl)-4-dicyclohexylphosphino-2-dicyclohexylphosphinomethylpyrrolidine (R-Cycapp) II (R₁ = Cy; R₂ = Ph, Me₃C, Cy) were prepd. by various synthetic methods. Rhodium(I) complexes of I and II showed high catalytic activity for hydrogenation of various kinds of prochiral ketones, which were reduced smoothly to the corresponding optically active hydroxy compds., under hydrogen at atm. pressure and ambient temp. The neutral rhodium(I) complexes (diphosphine-Rh) hydrogenated .alpha.-ketoamides and .alpha.-ketopantolactone in fairly high optical yields (66-77% ee). In the hydrogenation of N-(.alpha.-ketoacyl)-.alpha.-amino esters, the Cydiop-Rh catalyst showed a marked contrast to the diop-Rh system; in the hydrogenation of the Me ester of N-(phenylglyoxyl)-(S)-.alpha.-phenylalanine, 72% de was attained with little double asym. induction by the chiral center in the substrate.

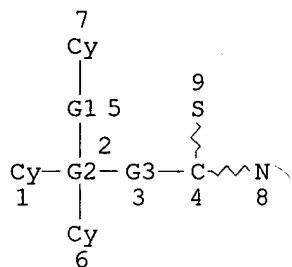
REFERENCE 2: 97:39003 Fully alkylated chiral diphosphines, RDIOP, and their rhodium(I) complexes. Tani, Kazuhide; Suwa, Kenichi; Yamagata, Tsuneaki; Otsuka, Sei (Fac. Eng. Sci., Osaka Univ., Osaka, 560, Japan). Chem. Lett. (3), 265-8 (English) 1982. CODEN: CMLTAG. ISSN: 0366-7022.

GI



AB Tetralkyl analogs of DIOP [2,3-O-isopropylidene-2,3-dihydroxy-1,4-bis(diphenylphosphino)butane], (-)-EtDIOP (I; R = Et), (-)-Me₂CHDIOP (I; R = Me₂CH), and (-)-CyDIOP (I; R = cyclohexyl), were prepd. by the reaction of (+)-2,3-O-isopropylidene-2,3-dihydroxy-1,4-difluorobutane and the corresponding LiPR₂. I form cationic Rh(I) complexes, [Rh((-)-RDIOP](NBD)]ClO₄ (NBD = norbornadiene, same RDIOP), which show remarkable reactivity toward H₂ in the hydrogenation of ketones.

=> d l36 que stat;d 1-2 ide cbib abs
L31 STR



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

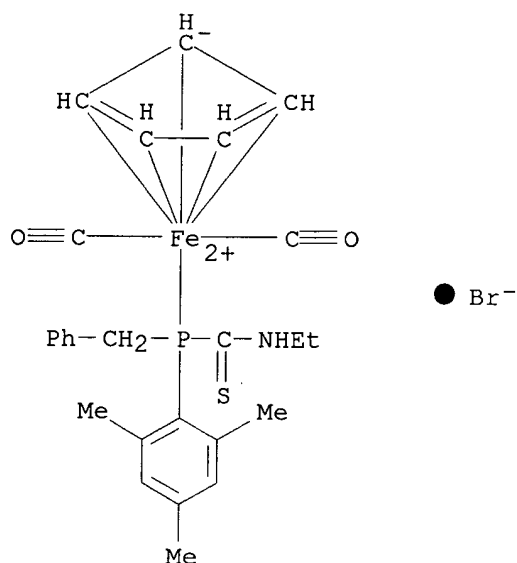
GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 9

STEREO ATTRIBUTES: NONE
 L36 2 SEA FILE=REGISTRY SSS FUL L31

100.0% PROCESSED 367681 ITERATIONS
 SEARCH TIME: 00.00.17

2 ANSWERS

L36 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2002 ACS
 RN 217633-00-6 REGISTRY
 CN Iron(1+), dicarbonyl(.eta.5-2,4-cyclopentadien-1-yl)[N-ethyl-1-(phenylmethyl)-1-(2,4,6-trimethylphenyl)phosphinecarbothioamide-.kappa.P]-, bromide (9CI) (CA INDEX NAME)
 MF C26 H29 Fe N O2 P S . Br
 CI CCS
 SR CA
 LC STN Files: CA, CAPLUS

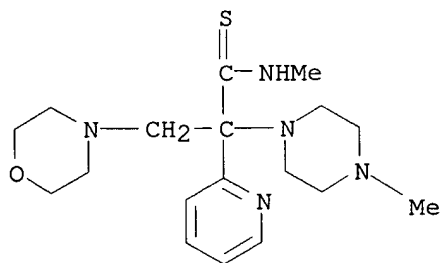


1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 130:66581 Transition metal-substituted phosphines, arsines, and stibines. Part 60. Ferrio(thiocarbamoyl)phosphines $\text{Cp}(\text{OC})_2\text{Fe}-\text{P}(\text{Mes})[\text{C}(:\text{S})-\text{N}(\text{R})\text{H}]$ ($\text{R} = \text{Me}, \text{Et}, \text{tBu}$). Buildup from the ferrio-mesitylphosphine $\text{Cp}(\text{OC})_2\text{Fe}-\text{P}(\text{Mes})\text{H}$ and organo isothiocyanates, quaternization with alkyl halides and oxidation with sulfur. Malisch, Wolfgang; Thirase, Katharina; Reising, Joachim (Institut Anorganische Chemie, Universitaet Wuerzburg, Wuerzburg, D-97074, Germany). Zeitschrift fuer Naturforschung, B: Chemical Sciences, 53(10), 1084-1091 (German) 1998. CODEN: ZNBSEN. ISSN: 0932-0776. Publisher: Verlag der Zeitschrift fuer Naturforschung.

AB The ferrio-phosphine $\text{Cp}(\text{OC})_2\text{Fe}-\text{P}(\text{Mes})\text{H}$, obtained by deprotonation of $\{\text{Cp}(\text{OC})_2[\text{H}_2(\text{Mes})\text{P}]\text{Fe}\}\text{BF}_4$ ($\text{Mes} = \text{mesityl}$), reacts with RNCS ($\text{R} = \text{Me}, \text{Et}, \text{Ph}$) to give the corresponding ferrio-phosphines $\text{Cp}(\text{OC})_2\text{Fe}-\text{P}(\text{Mes})[\text{C}(:\text{S})-\text{N}(\text{R})\text{H}]$ (same R , 6a-c). Quaternization of 6 ($\text{R} = \text{Me}, \text{Et}$) at the P atom with alkyl halides R_1X ($\text{R}_1 = \text{Me}$ or Et , $\text{X} = \text{I}$; $\text{R}_1 = \text{PhCH}_2$, $\text{X} = \text{Br}$) yields $\{\text{Cp}(\text{OC})_2\text{Fe}-\text{P}(\text{Mes})(\text{R}_1)[\text{C}(:\text{S})-\text{N}(\text{H})(\text{R})]\}\text{X}$ (8; $\text{R} = \text{R}_1 = \text{Me}$, $\text{X} = \text{I}$; $\text{R} = \text{Et}$, $\text{R}_1 = \text{Me}$, $\text{X} = \text{I}$; $\text{R} = \text{Et}$, $\text{R}_1 = \text{PhCH}_2$, $\text{X} = \text{Br}$), whereas oxidn. with elemental S affords the ferrio-thiophosphoranes $\text{Cp}(\text{OC})_2\text{Fe}-\text{P}(:\text{S})(\text{Mes})[\text{C}(:\text{S})-\text{N}(\text{R})\text{H}]$ (10; $\text{R} = \text{Me}, \text{Et}$). 10B ($\text{R} = \text{Et}$) is alkylated with MeI to give $\{\text{Cp}(\text{OC})_2\text{Fe}-\text{P}(\text{SMe})(\text{Mes})[\text{C}(:\text{S})-\text{N}(\text{Et})\text{H}]\}\text{I}$. The structure of 8b ($\text{R} = \text{Et}$, $\text{R}_1 = \text{Me}$, $\text{X} = \text{I}$) was detd. by x-ray anal.

L36 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2002 ACS
RN 59769-18-5 REGISTRY
CN 4-Morpholinepropanethioamide, N-methyl-.alpha.-(4-methyl-1-piperazinyl)-.alpha.-2-pyridinyl- (9CI) (CA INDEX NAME)
FS 3D CONCORD
MF C18 H29 N5 O S
LC STN Files: CA, CAPLUS, USPATFULL



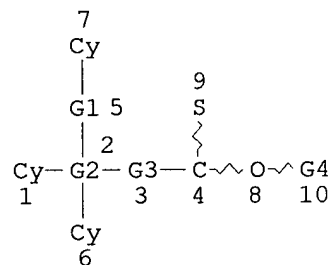
PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 85:46700 2-Alkoxy(and 2-amino)-3-amino-2-heterocyclic-thiopropenamides. Loev, Bernard (Smithkline Corp., USA). U.S. US 3948892 19760406, 10 pp. Division of U.S. 3,860,592. (English). CODEN: USXXAM. APPLICATION: US 1974-514684 19741015.

AB RR1(MeO)CSNHR2 (I, R = H, morpholinomethyl; R1 = 2-pyridyl, 2-pyrrolyl; R2 = H, Me, cyclohexyl, allyl, Ph etc.) (.apprx.20 compds.) were prepd. Thus, 2-(chloromethyl)pyridine was treated with MeONa in MeOH to give 2-(methoxymethyl)pyridine which was then treated with MeNCS in presence of PhLi to give I (R = R2 = H, R1 = 2-pyridyl). Refluxing the last with morpholine and formaldehyde for 48 hr gave I (R = morpholinomethyl, R1 = 2-pyridyl, R2 = H). I (R = morpholinomethyl) inhibits gastric acid secretion in pylorus-ligated rats at 10-50 mg/kg orally.

=> d 139 que stat
L37 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
VAR G4=S/O
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

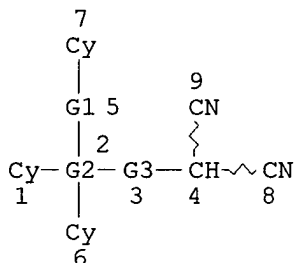
GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 10

STEREO ATTRIBUTES: NONE
L39 0 SEA FILE=REGISTRY SSS FUL L37

100.0% PROCESSED 41529 ITERATIONS
 SEARCH TIME: 00.00.06

0 ANSWERS

=> d l42 que stat;d 1-3 ide cbib abs
 L40 STR



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

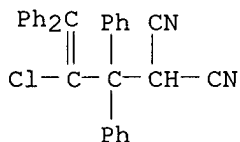
GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 9

STEREO ATTRIBUTES: NONE
 L42 3 SEA FILE=REGISTRY SSS FUL L40

100.0% PROCESSED 68859 ITERATIONS
 SEARCH TIME: 00.00.07

3 ANSWERS

L42 ANSWER 1 OF 3 REGISTRY COPYRIGHT 2002 ACS
 RN 19937-41-8 REGISTRY
 CN Malononitrile, (2-chloro-1,1,3,3-tetraphenylallyl)- (8CI) (CA INDEX NAME)
 MF C30 H21 Cl N2
 LC STN Files: BEILSTEIN*, CA, CAPLUS
 (*File contains numerically searchable property data)



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

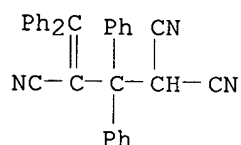
1 REFERENCES IN FILE CA (1962 TO DATE)
 1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

Searched by: Mary Hale 308-4258 CM-1 1E01

REFERENCE 1: 69:59302 Stable carbenoids. XXXI. .alpha.-Functional vinylolithium compounds from .alpha.-haloacrylic acid derivatives. Koebrich, Gert; Trapp, Horst; Akhtar, Ali (Univ. Heidelberg, Heidelberg, Ger.). Chem. Ber., 101(8), 2644-59 (German) 1968. CODEN: CHBEAM.

AB .alpha.-Haloacrylic acids R:C(Cl)CN and R:C(Br)CO2R1 (derived from carbenoids) were treated with organolithium compds. in tetrahydrofuran at low temps. to give R:C(CN-)Li+ and R:C-(CO2R1)Li+. The halogen-metal exchange takes place very quickly and overrides addn. reactions, such as to the CN or CO2R1 groups. It causes anomalous results in the cyanation and carboxylation of R:C-(Br)Li and may compete with the deprotonation of the carboxyl group. 27 references.

L42 ANSWER 2 OF 3 REGISTRY COPYRIGHT 2002 ACS
 RN 19937-39-4 REGISTRY
 CN 3-Butene-1,1,3-tricarbonitrile, 2,2,4,4-tetraphenyl- (8CI) (CA INDEX NAME)
 MF C31 H21 N3
 LC STN Files: BEILSTEIN*, CA, CAPLUS
 (*File contains numerically searchable property data)



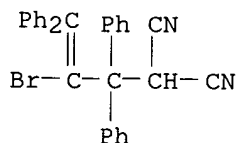
PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
 1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 69:59302 Stable carbenoids. XXXI. .alpha.-Functional vinylolithium compounds from .alpha.-haloacrylic acid derivatives. Koebrich, Gert; Trapp, Horst; Akhtar, Ali (Univ. Heidelberg, Heidelberg, Ger.). Chem. Ber., 101(8), 2644-59 (German) 1968. CODEN: CHBEAM.

AB .alpha.-Haloacrylic acids R:C(Cl)CN and R:C(Br)CO2R1 (derived from carbenoids) were treated with organolithium compds. in tetrahydrofuran at low temps. to give R:C(CN-)Li+ and R:C-(CO2R1)Li+. The halogen-metal exchange takes place very quickly and overrides addn. reactions, such as to the CN or CO2R1 groups. It causes anomalous results in the cyanation and carboxylation of R:C-(Br)Li and may compete with the deprotonation of the carboxyl group. 27 references.

L42 ANSWER 3 OF 3 REGISTRY COPYRIGHT 2002 ACS
 RN 19937-38-3 REGISTRY
 CN Malononitrile, (2-bromo-1,1,3,3-tetraphenylallyl)- (8CI) (CA INDEX NAME)
 MF C30 H21 Br N2
 LC STN Files: BEILSTEIN*, CA, CAPLUS
 (*File contains numerically searchable property data)

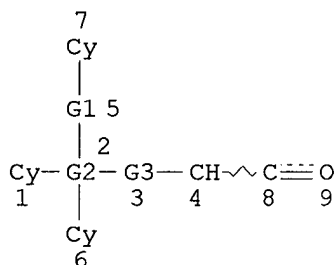


PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 69:59302 Stable carbenoids. XXXI. .alpha.-Functional vinylolithium compounds from .alpha.-haloacrylic acid derivatives. Koebrich, Gert; Trapp, Horst; Akhtar, Ali (Univ. Heidelberg, Heidelberg, Ger.). Chem. Ber., 101(8), 2644-59 (German) 1968. CODEN: CHBEAM.
AB .alpha.-Haloacrylic acids R:C(Cl)CN and R:C(Br)CO2R1 (derived from carbenoids) were treated with organolithium compds. in tetrahydrofuran at low temps. to give R:C(CN-)Li+ and R:C-(CO2R1)Li+. The halogen-metal exchange takes place very quickly and overrides addn. reactions, such as to the CN or CO2R1 groups. It causes anomalous results in the cyanation and carboxylation of R:C-(Br)Li and may compete with the deprotonation of the carboxyl group. 27 references.

=> d l44 que stat
L43 STR



REP G1=(1-3) C
VAR G2=C/N/P/SI/GE
REP G3=(0-4) CH2
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 9

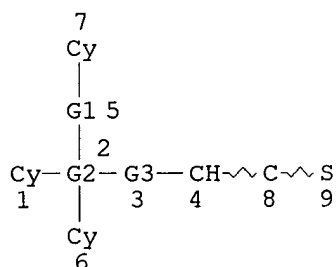
STEREO ATTRIBUTES: NONE
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SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
BATCH **INCOMPLETE**
PROJECTED ITERATIONS: EXCEEDS 1000000
PROJECTED ANSWERS: EXCEEDS 0

=> d l46 que stat
L45 STR

Searched by: Mary Hale 308-4258 CM-1 1E01



REP G1=(1-3) C
 VAR G2=C/N/P/SI/GE
 REP G3=(0-4) CH2
 NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 9

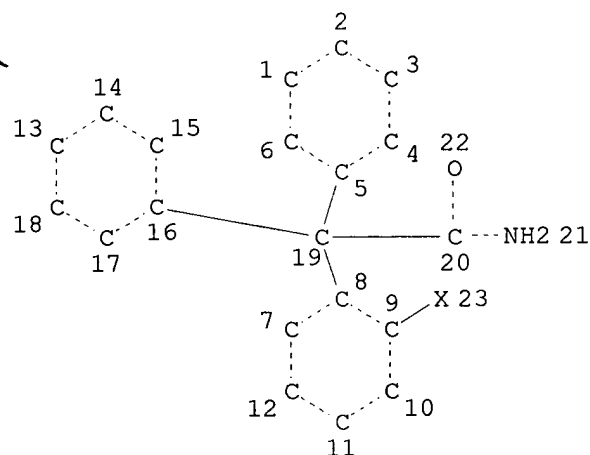
STEREO ATTRIBUTES: NONE
 L46 0 SEA FILE=REGISTRY SSS SAM L45

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 INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
 SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**
 BATCH **INCOMPLETE**
 PROJECTED ITERATIONS: EXCEEDS 1000000
 PROJECTED ANSWERS: EXCEEDS 0

=> d l49 que stat;d 1-10 ide cbib abs
 L47 STR



NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

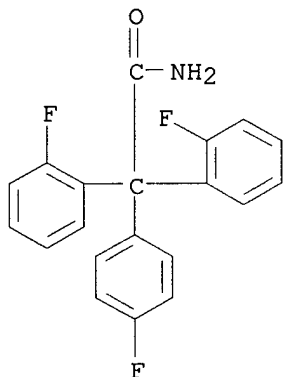
Searched by: Mary Hale 308-4258 CM-1 1E01

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 23

STEREO ATTRIBUTES: NONE
L49 10 SEA FILE=REGISTRY SSS FUL L47

100.0% PROCESSED 252 ITERATIONS 10 ANSWERS
SEARCH TIME: 00.00.01

L49 ANSWER 1 OF 10 REGISTRY COPYRIGHT 2002 ACS
RN 289656-69-5 REGISTRY
CN Benzeneacetamide, 2-fluoro-.alpha.-(2-fluorophenyl)-.alpha.-(4-
fluorophenyl)- (9CI) (CA INDEX NAME)
MF C20 H14 F3 N O
SR CA
LC STN Files: CA, CAPLUS, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

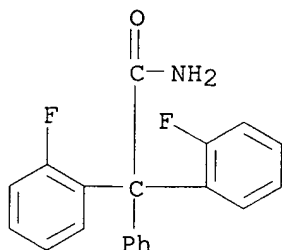
REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo

Searched by: Mary Hale 308-4258 CM-1 1E01

half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

L49 ANSWER 2 OF 10 REGISTRY COPYRIGHT 2002 ACS
RN 289656-67-3 REGISTRY
CN Benzeneacetamide, 2-fluoro-.alpha.-(2-fluorophenyl)-.alpha.-phenyl- (9CI)
(CA INDEX NAME)
MF C20 H15 F2 N O
SR CA
LC STN Files: CA, CAPLUS, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

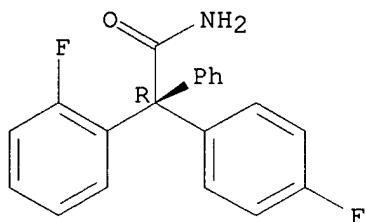
REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

L49 ANSWER 3 OF 10 REGISTRY COPYRIGHT 2002 ACS
RN 289656-63-9 REGISTRY
CN Benzeneacetamide, 2-fluoro-.alpha.-(4-fluorophenyl)-.alpha.-phenyl-,
(.alpha.R)- (9CI) (CA INDEX NAME)
FS STEREOSEARCH
MF C20 H15 F2 N O
SR CA
LC STN Files: CA, CAPLUS, USPATFULL

Searched by: Mary Hale 308-4258 CM-1 1E01

Absolute stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

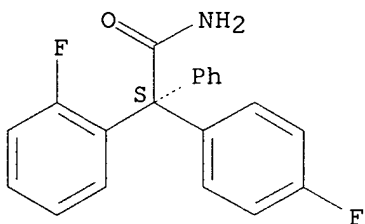
1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

L49 ANSWER 4 OF 10 REGISTRY COPYRIGHT 2002 ACS
RN 289656-61-7 REGISTRY
CN Benzeneacetamide, 2-fluoro-.alpha.-(4-fluorophenyl)-.alpha.-phenyl-, (.alpha.S)- (9CI) (CA INDEX NAME)
FS STEREOSEARCH
MF C20 H15 F2 N O
SR CA
LC STN Files: CA, CAPLUS, USPATFULL

Absolute stereochemistry.



Searched by: Mary Hale 308-4258 CM-1 1E01

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degradn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

L49 ANSWER 5 OF 10 REGISTRY COPYRIGHT 2002 ACS

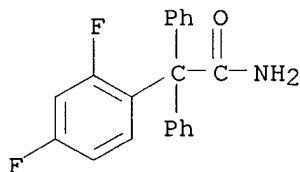
RN 289656-55-9 REGISTRY

CN Benzeneacetamide, 2,4-difluoro-.alpha.,.alpha.-diphenyl- (9CI) (CA INDEX NAME)

MF C20 H15 F2 N O

SR CA

LC STN Files: CA, CAPLUS, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601

Searched by: Mary Hale 308-4258 CM-1 1E01

19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

L49 ANSWER 6 OF 10 REGISTRY COPYRIGHT 2002 ACS

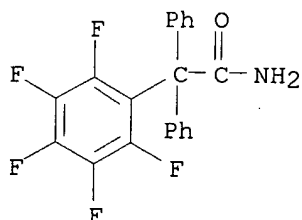
RN 289656-51-5 REGISTRY

CN Benzeneacetamide, 2,3,4,5,6-pentafluoro-.alpha.,.alpha.-diphenyl- (9CI)
(CA INDEX NAME)

MF C20 H12 F5 N O

SR CA

LC STN Files: CA, CAPLUS, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

L49 ANSWER 7 OF 10 REGISTRY COPYRIGHT 2002 ACS

RN 289656-49-1 REGISTRY

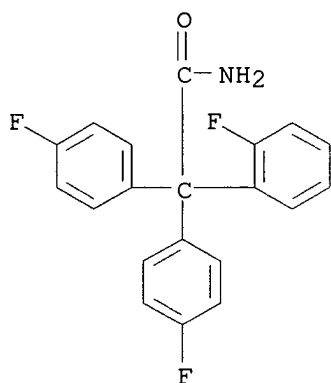
CN Benzeneacetamide, 2-fluoro-.alpha.,.alpha.-bis(4-fluorophenyl)- (9CI) (CA INDEX NAME)

MF C20 H14 F3 N O

SR CA

LC STN Files: CA, CAPLUS, USPATFULL

Searched by: Mary Hale 308-4258 CM-1 1E01



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

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L49 ANSWER 8 OF 10 REGISTRY COPYRIGHT 2002 ACS

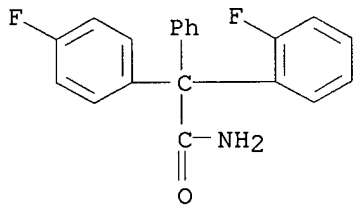
RN 289656-42-4 REGISTRY

CN Benzeneacetamide, 2-fluoro-.alpha.-(4-fluorophenyl)-.alpha.-phenyl- (9CI)
(CA INDEX NAME)

MF C20 H15 F2 N O

SR CA

LC STN Files: CA, CAPLUS, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:187954 Gardos channel antagonists, their preparation, and their therapeutic use. McNaughton-Smith, Grant Andrew; Rigdon, Gregory Cooksey; Stocker, Jonathan Walter (Icagen, Inc., USA). PCT Int. Appl. WO 2000050026 A1 20000831, 53 pp. DESIGNATED STATES: W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-US3663 20000210. PRIORITY: US 1999-PV135511 19990223; US 1999-386601 19990831.

AB Inhibitors of potassium flux are disclosed. The inhibitors show surprising resistance to degrdn. in biol. media and enhanced in vivo half-lives relative to non-fluorine substituted homologues. Methods for the use of these compds. include treating sickle cell disease, preventing erythrocyte dehydration, and inhibiting potassium flux. Compds. of the invention include fluorinated triphenylacetamides (prepn. described).

L49 ANSWER 9 OF 10 REGISTRY COPYRIGHT 2002 ACS

RN 197526-18-4 REGISTRY

CN Benzeneacetamide, 2-chloro-.alpha.,.alpha.-diphenyl- (9CI) (CA INDEX NAME)

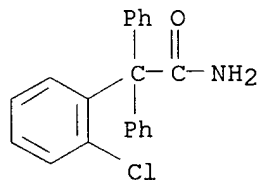
OTHER NAMES:

CN (2-Chlorophenyl)diphenylacetamide

MF C20 H16 Cl N O

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1962 TO DATE)

3 REFERENCES IN FILE CAPLUS (1962 TO DATE)

Searched by: Mary Hale 308-4258 CM-1 1E01